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### INTERNATIONAL SEARCH REPORT

Internameal Application No

PCT/US2004/031224 A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C12P21/02 C12N9/06 C12N15/53 According to International Patent Classification (IPC) or to both national classification and IPC Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12N C12P B. FIELDS SEARCHED C12N C12P Documentation searched other than minimum documentation to the externt that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, Sequence Search, WPI Data, PAJ, EMBASE, BIOSIS C. DOCUMENTS CONSIDERED TO BE RELEVANT Category Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Α BOGOSIAN G ET AL: "Biosynthesis and 1-18, incorporation into protein of norleucine 27-42 by Escherichia coli." THE JOURNAL OF BIOLOGICAL CHEMISTRY. 5 JAN vol. 264, no. 1, 5 January 1989 (1989-01-05), pages 531-539, XP002324310 ISSN: 0021-9258 cited in the application the whole document A US 5 698 418 A (BRUNNER ET AL) 1-18. 16 December 1997 (1997-12-16) 27-42 cited in the application abstract -/--X Further documents are listed in the continuation of box C. X Patent family members are listed in annex. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance invention "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to Involve an Inventive step when the document is combined with one or more other such docu-"O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled in the art. "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 11 -07- 2005 15 April 2005

Authorized officer

Wiame, I

European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

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## INTERNATIONAL SEARCH REPORT

Intermental Application No
PCT/US2004/031224

C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	10170320047031224	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to daim No.	
A	JAKUBOWSKI H ET AL: "Editing of errors in selection of amino acids for protein synthesis." MICROBIOLOGICAL REVIEWS. SEP 1992, vol. 56, no. 3, September 1992 (1992-09), pages 412-429, XP002324311 ISSN: 0146-0749 page 425, column 1, last paragraph page 420, column 2, last paragraph - page 421, column 2, last paragraph	1-18, 27-42	
A	KATAOKA KUNISHIGE ET AL: "Alteration of substrate specificity of leucine dehydrogenase by site-directed mutagenesis."  JOURNAL OF MOLECULAR CATALYSIS B ENZYMATIC, vol. 23, no. 2-6, 1 September 2003 (2003-09-01), pages 299-309, XP002324312 ISSN: 1381-1177 the whole document	1-7, 16-18, 27-34, 36,38-40	
A	US 5 789 234 A (BERTELSEN ET AL) 4 August 1998 (1998-08-04)	3-7, 16-18, 30,32, 33,37-40	

## **INTERNATIONAL SEARCH REPORT**

International application No. PCT/US2004/031224

Box II Observations where certain claims were found unsearchable (Continuation of Item 2 of first sheet)							
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:							
Claims Nos.:     because they relate to subject matter not required to be searched by this Authority, namely:							
Claims Nos.:     because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:							
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).							
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)							
This International Searching Authority found multiple Inventions in this International application, as follows:							
see additional sheet							
As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.							
As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.							
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:							
4. X No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  1-18, 27-42 (all completely)							
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.							

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-18, 27-42 (all completely)

use of non-standard amino acid degrading protein in a method for reducing the incorporation of non-standard amino acids into a heterologous protein expressed by a microorganism and in methods of isolating a heterologous protein

2. claims: 19-26 (all completely)

E. coli glutamate dehydrogenase protein comprising a L92K variation, DNA molecule encoding said protein and E. coli cell comprising said DNA molecule or expressing said protein

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## INTERNATIONAL SEARCH REPORT

Information on patent family members

Internamial Application No
PCT/US2004/031224

		101/03200.7032221		
Patent document cited in search report	Publication date		Patent family member(s)	Publication date
US 5698418 A	16-12-1997	US AT AU DE DE DK EP JP JP WO	5622845 A 152176 T 3058489 A 68927997 D1 68927997 T2 181790 A 0401254 A1 0737752 A2 2879063 B2 3502758 T 8907651 A2	22-04-1997 15-05-1997 06-09-1989 28-05-1997 11-09-1997 30-07-1990 12-12-1990 16-10-1996 05-04-1999 27-06-1991 24-08-1989
US 5789234 A	04-08-1998	USTA CE THE JUNE AND THE STATE OF THE STATE	6319685 B1 283922 T 2009306 A1 69034179 D1 382403 T3 0382403 A2 110517 B1 54414 A2 2072995 C 5041982 A 7071498 B 157983 B1 1829 A 900531 A 2129606 C1 198355 T 629795 B2 2057688 A 1341454 C 3856446 D1 3856446 D1 3856446 T2 448988 A 0308067 A2 0529742 A1 883722 A 935385 A 213227 B1 212071 B3 9500552 A3 2104281 A 2979172 B2 136611 B1 1656 A 883592 A 2080385 C1	20-11-2001 15-12-2004 06-08-1990 05-01-2005 21-03-2005 16-08-1990 14-02-2003 28-02-1991 25-07-1996 23-02-1993 02-08-1995 16-11-1998 25-08-1995 07-08-1990 27-04-1999 15-01-2001 15-10-1992 16-02-1989 20-04-2004 01-02-2001 12-07-2001 12-07-2001 12-07-2001 15-02-1989 22-03-1989 03-03-1993 15-02-1989 01-12-1993 28-07-1997 28-02-1996 30-10-1995 17-04-1990 15-11-1999 25-04-1998 25-07-1995 15-02-1989 27-05-1997